=> fil hcaplu FILE 'HCAPLUS' ENTERED AT 09:28:13 ON 24 JUN 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 24 Jun 2003 VOL 138 ISS 26 FILE LAST UPDATED: 23 Jun 2003 (20030623/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d stat que 1 SEA FILE=REGISTRY ABB=ON COBALT/CN L4343310 SEA FILE=HCAPLUS ABB=ON L4 OR COBALT L8 191148 SEA FILE=HCAPLUS ABB=ON TRANSITION(W)METAL? OR LANTHANIDE? Ь9 10 SEA FILE=HCAPLUS ABB=ON (L8 OR L9)(L)(CHELAT? OR COMPLEX?) T.11 AND (BACTER? OR FUNG?)(W)(DETECT? OR IDEN? OR ASSAY?)

=> d ibib abs hitrn 111 1-10

L11 ANSWER 1 OF 10 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER:

2001:863434 HCAPLUS

DOCUMENT NUMBER:

136:2484

TITLE:

Mass spectrometric detection of polypeptides

INVENTOR(S):

Little, Daniel; Koster, Hubert; Higgins, G. Scott;

Lough, David

PATENT ASSIGNEE(S):

Sequenom, Inc., USA

SOURCE:

U.S., 50 pp., Cont.-in-part of U.S. Ser. No. 922,201.

CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICA	
US 6322970 B1 20011127 US 1998 US 6207370 B1 20010327 US 1997 EP 1296143 A2 20030326 EP 2002 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, I IE, SI, LT, LV, FI, RO, MK, CY, AL US 6387628 B1 20020514 US 2000 US 2003003465 A1 20030102 US 2001 PRIORITY APPLN. INFO.: US 1997-92 EP 1998-94	-922201 19970902 -25544 19980902 T, LI, LU, NL, SE, MC, PT, -664977 20000918 -7557 20011106 2201 A2 19970902

A3 19980902 US 1998-146054 US 2000-664977 A1 .20000918

A process for detg. the identity of a target polypeptide using mass AΒ spectroscopy is provided. Depending on the target polypeptide to be identified, a process as disclosed can be used, for example, to diagnose a genetic disease or chromosomal abnormality, a predisposition to a disease or condition, or infection by a pathogenic organism; or for detg. identity or heredity. Kits for performing the disclosed processes also are provided. A process for obtaining information on a sequence of a target nucleic acid mol. by detg. the identity of a polypeptide encoded by the nucleic acid mol. comprises: (a) prepg. the encoded polypeptide from a target nucleic acid mol. by in vitro translation, or by in vitro transcription followed by translation, of the target nucleic acid mol.; (b) detg. the mol. mass of the encoded polypeptide by mass spectrometry; and (c) detg. the identity of the polypeptide by comparing the mol. mass of the polypeptide with the mol. mass of a corresponding known polypeptide, thereby obtaining information on a sequence of nucleotides in the target nucleic acid mol.

7440-48-4D, Cobalt, ions, supported chelates, IT

uses

RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (in isolation of encoded tagged polypeptide; mass spectrometric detection of polypeptides)

REFERENCE COUNT:

THERE ARE 269 CITED REFERENCES AVAILABLE FOR 269 THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

L11 ANSWER 2 OF 10 HCAPLUS COPYRIGHT 2003 ACS 2001:331247 HCAPLUS

ACCESSION NUMBER:

134:337915

DOCUMENT NUMBER: TITLE:

Competitive apo-peroxidase assay

INVENTOR(S):

Pugia, Michael J.

PATENT ASSIGNEE(S):

Bayer Corporation, USA

SOURCE:

U.S., 8 pp., Cont.-in-part of U.S. Ser. No. 990,389,

abandoned. CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 6228602	В1	20010508	US 1999-415491	19991012
AU 9897074	A1	19990701	AU 1998-97074	19981211
AU 742373	В2	20020103		
JP 11237382	A2	19990831	JP 1998-354203	19981214
PRIORITY APPLN. INFO.:	:	•	US 1997-990389 B2	19971215

Disclosed is an assay for an analyte in a fluid test sample such as urine which involves combining the fluid test sample with a reagent system comprising an apo-peroxidase, a redox dye, a peroxide and a metal porphyrin which is bound to an analyte/analyte specific binding partner complex, which complex has a combined mol. wt. of at least about 180 K Daltons. When this conjugate interacts with analyte in the fluid test sample, a portion of the specific binding partner is dissocd. from the complex thereby enabling the metal porphyrin to reconstitute with the apo-peroxidase. The reconstituted peroxidase can interact with the peroxide and redox dye to provide a colored response to analyte in the fluid test sample. An ascorbate and Hb-resistant reagent for detecting

Lucas 10 /082,618

peroxidase activity was combined with apo-Hb, Fe hematin-LPS conjugate, and antibody to anti-rabbit bacterial LPS in one reagent. The complete reagent detected bacterial cells in urine with three species of gram neg. cells being detected.

IT 7440-48-4D, Cobalt, deuteroporphyrin complexes

, biological studies

RL: ARG (Analytical reagent use); BPR (Biological process); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study); PROC (Process); USES (Uses)

(competitive apo-peroxidase assay)

REFERENCE COUNT:

THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L11 ANSWER 3 OF 10 HCAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 2001:284225 HCAPLUS

ACCESSION NUMBER: DOCUMENT NUMBER:

134:307594

TITLE:

Antibiotic-metal complexes in the detection of

gram-negative bacteria and other biological analytes

INVENTOR(S):

Olstein, Alan D.; Feirtag, Joellen M.

PATENT ASSIGNEE(S):

USA

SOURCE:

PCT Int. Appl., 30 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

	PATENT NO.	KIND D	ATE	APPLICATION NO. DATE
	WO 2001027628 W: AE, AG, CR, CU, HU, ID, LU, LV, SD, SE,	A1 2 AL, AM, CZ, DE, IL, IN, MA, MD, SG, SI,	0010419 AT, AU, AZ DK, DM, DZ IS, JP, KE MG, MK, MN SK, SL, TJ	WO 2000-US28577 20001013 Z, BA, BB, BG, BR, BY, BZ, CA, CH, CN, Z, EE, ES, FI, GB, GD, GE, GH, GM, HR, E, KG, KP, KR, KZ, LC, LK, LR, LS, LT, N, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, J, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, Z, MD, RU, TJ, TM
		**** * T O	MOTOR MOTOR	n et en un un nom Ai, de, en ei
	DE, DK, CF, CG,	ES, FI, CI, CM,	FR, GB, GF GA, GN, GW	W, ML, MR, NE, SN, TD, TG US 1999-159142P P 19991013
	RITY APPLN. INFO		and motal	le are provided that are useful
AB	in detecting ba useful in detection preferably chel is a polymyxin, thereof. Metho Polymyxin B-cob the cell titrn.	cteria an ting gram ated comp a colist ds of usi alt compl of Esche	nd other bin neg. bact blexes when tin, an aming the corlex (preprietable collection col	teria. The complexes are trein the antibiotic tinoglycoside, or an analog or fragment timplexes are also provided. In given) was used in the complexes are also provided. In O157:H7. Chemiluminescence was
IT	7440-48-4DP, Co preparation RL: ARG (Analyt preparation); F (antibiotic-	cical read ANST (Anal-metal con	mplex with gent use); lytical st mplexes in iol. analy	PRP (Properties); SPN (Synthetic cudy); PREP (Preparation); USES (Uses) detection of gram-neg.
ΙT	7440-48-4D, Col chelates with a			, complexes of gical studies , тип (Therapeutic use); ANST (Analytica
				, while congradeditic doc// force (thice)

T7440-48-4D, Cobalt, and isotopes, complexes or
chelates with antibiotics, biological studies
RL: ARG (Analytical reagent use); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)

(antibiotic-metal complexes in detection of gram-neg.

bacteria and other biol. analytes) THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT REFERENCE COUNT: - 2

L11 ANSWER 4 OF 10 HCAPLUS COPYRIGHT 2003 ACS 1999:404774 HCAPLUS

ACCESSION NUMBER:

DOCUMENT NUMBER:

131:41802

TITLE:

Assay system utilizing apo-peroxidase, a

hydroperoxide, a redox dye and a metal porphyrin which

is bound to analyte or analyte-specific binding

partner conjugate Pugia, Michael J.

INVENTOR(S): PATENT ASSIGNEE(S):

Bayer Corp., USA Eur. Pat. Appl., 14 pp.

SOURCE:

CODEN: EPXXDW

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

APPLICATION NO. DATE KIND DATE _____ PATENT NO. -----EP 1998-122881 19981202 19990623 EP 924521 A1 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO EP 924521 AU 1998-97074 19990701 A1 AU 9897074 20020103 В2 AU 742373 19981214 JP 1998-354203 19990831 A2 JP 11237382 US 1997-990389 A 19971215

Disclosed is an assay for an analyte in a fluid test sample such as urine PRIORITY APPLN. INFO.: which involves combining the fluid test sample with a reagent system comprising an apo-peroxidase, a redox dye, a hydroperoxide and a metal porphyrin which is bound to an analyte/analyte specific binding partner which conjugate has a combined mol. wt. of at least about 180 K Daltons. When this conjugate interacts with analyte in the fluid test sample, a portion of the specific binding partner is dissocd. from the conjugate thereby enabling the metal porphyrin to reconstitute with the apo-peroxidase. The reconstituted peroxidase can interact with the hydroperoxide and redox dye to provide a colored response to analyte in the fluid test sample.

7440-48-4D, Cobalt, complex with IT

RL: ARU (Analytical role, unclassified); BPR (Biological process); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study); PROC (Process)

(assay system utilizing apo-peroxidase, a hydroperoxide, a redox dye and a metal porphyrin which is bound to analyte or analyte-specific binding partner conjugate)

REFERENCE COUNT:

THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT 5

L11 ANSWER 5 OF 10 HCAPLUS COPYRIGHT 2003 ACS 1994:172315 HCAPLUS

ACCESSION NUMBER: DOCUMENT NUMBER:

TITLE:

Microbial degradation of metal complexed cyanides and

thiocyanate from mining wastewaters

AUTHOR (S):

Boucabeille, C.; Bories, A.; Ollivier, P.; Michel, G.

CORPORATE SOURCE:

Lab. Biotechnol. Environ., INRA, Narbonne, 11100, Fr. Environmental Pollution (Oxford, United Kingdom)

(1994), 84(1), 59-67

CODEN: ENPOEK; ISSN: 0269-7491

DOCUMENT TYPE:

SOURCE:

Journal English

LANGUAGE: A microbiol. examn. of the soil from a CN- wastewater storage basin was carried out. The storage basin contained water from the cyanidation process of Au extn., and it was composed principally of CN-, metal complexed cyanide, mainly cuprocyanide, ferro-ferricyanides and thiocyanate. Pseudomonas species were the principal bacteria identified in the soil. Using the storage basin soil as a seed sludge, its potential for the biodegrdn. of all the cyanide complexes in the mining wastewater was studied in the lab., using batch, fed-batch and continuous processes. The NH3 and SO42- produced were quantified. In the continuous process, total degrdn. of all cyanide was obsd. at a diln. rate of 0.066/day.

L11 ANSWER 6 OF 10 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER:

1992:17788 HCAPLUS

DOCUMENT NUMBER:

116:17788

TITLE:

Enzyme amplified lanthanide chelate

luminescence assay

INVENTOR(S):

Evangelista, Ramon A.; Templeton, Eva F. Gudgin;

Pollak, Alfred

PATENT ASSIGNEE(S):

Kronem Systems Inc., Can. PCT Int. Appl., 103 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PAT	CENT 1	NO.		KI	ND	DATE			A	PPLI	CATI	ои ис). 	DATE		•	•
WO	9108 W:	AΤ.	AU,	BB.	BG,	BR,	CA,	CH,	DE,	DK,	ES,	FI,	GB,	1990: GR,	1113 HU,	JP,	KP,
	RW:	AT,	LK, BE, ML,	BF,	ВJ,	CF,	CG,	CH,	CM,	RO, DE,	SD, DK,	ES,	FR,	GA,	GB,	GR,	IT,
	2067	394	mu,	A.	Α.	1991	0605	,	С	A 19	90-2	0673	94	1990	1113		
AU	2067 9067	150		A.	1	1991	0626			U 19 P 19				1990 1990			
	5446 5446	62		В.	1	1996	0609 0918										
• •	5262	299	BE,			1993	1116 1015		Ū	S 19	90-6	121/	Ŧ	NL, 1990 1990	1112		
PRIORIT	1431 Y APP		INFO	_		1990	1015		GB 1		2750	3	Α	1989	1204		

MARPAT 116:17788 OTHER SOURCE(S):

A method and compds. useful for this method of enzyme-amplified signal detection in anal. assays requiring extremely high sensitivity comprises use of a substrate which is capable of being transformed by an enzyme from a compd. which does not form a luminescent lanthanide chelate into a product which does. This method is particularly useful in time-resolved luminescence anal. Thus, salicylaldehyde was oxidized by xanthine oxidase to salicylic acid. The reaction was stopped by adding EDTA, TbCl3, and NaOH and the fluorescence of the 1:1:1

> EDTA-salicylic acid-Tb complex was detd. at 545 nm. detection limit was .apprx.5 .times. 10-6 U/mL, a sensitivity comparable to radioactive methods. Substrates for esterases, .beta.-galactosidase, alk. phosphatase, and glucose oxidase were synthesized and the assay using these compds. were demonstrated.

L11 ANSWER 7 OF 10 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER:

1990:494380 HCAPLUS

DOCUMENT NUMBER:

113:94380

TITLE:

Method for minimizing interference by reductants when

detecting cells in biological fluids

INVENTOR(S):

Belly, Robert T.; Sullivan, Sheryl S.; Schmittou, Eric

PATENT ASSIGNEE(S):

Eastman Kodak Co., USA

SOURCE:

U.S., 6 pp. CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

APPLICATION NO. DATE KIND DATE PATENT NO. -----_____ ----19870611 US 1987-60559 19900327 Α tis 4912035 19870611

US 1987-60559 PRIORITY APPLN. INFO.: A method including cell sepn. (by filtration, centrifugation, or pptn.), cell washing (with Fe2+-chelate soln. and nonionic surfactant soln.), and cell detection [by using a reducible dye, preferably a Co(III) complex] is adapted to minimize interference by reductants during cell (including bacteria, yeast, fungi, etc.) detection in biol. fluids (urine, blood, etc.). Thus, a urine sample was filtered through a cellulose acetate filter, the filter was then washed with Fe3+-EDTA and Triton X-100, and was removed for cell detection at 610 nm in the presence of Co3+ and 2,3-dimethoxy-5-methyl-1,4-benzoquinone. A cobalt chem. coating contg. (2,2'-bipyridine)bis(1,2-diaminoethane)cobalt (III) chloride, glucose, diammonium 2-[(5-carboxy-2-pyridyl)azo]-1naphthol-4-sulfonate, etc. was prepd. for cell detection. Several Fe3+chelates were also prepd. and tested; Fe3+-nitriloacetic acid, Fe3+-iminodiacetic acid, and Fe3+-N-methyliminodiacetic acid were very effective as washing reagents.

7440-48-4D, Cobalt, complexes TT

RL: ANST (Analytical study)

(cell detection with redox reagent contg. dye and)

L11 ANSWER 8 OF 10 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER:

1988:201342 HCAPLUS

DOCUMENT NUMBER:

108:201342

TITLE:

Analyte detection by means of fluorescent energy

transfer

INVENTOR(S):

Stavrianopoulos, Jannis; Rabbani, Elazar; Abrams,

Samuel B.; Wetmur, James Gerard

PATENT ASSIGNEE(S):

Enzo Biochem, Inc., USA Eur. Pat. Appl., 66 pp.

SOURCE:

CODEN: EPXXDW

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

Lucas 10 /082,618

PATENT NO	KIND	DATE	APPLICATION NO.	DATE
EP 242527 EP 242527	A2 A3 B1	19871028 19890614	EP 1987-102315	19870218
R: C. US 486810 CA 128533 JP 622408 PRIORITY APPLN AB An analyt energy em 2nd fluor E1 is abs fluoresce period. fluoresce Either E1	H, DE, FR, GB A A A A A A A A A A A A A A A A A A A	, GR, IT, L1 19890919 19910625 19871021 by binding the emitter (Exwhich is poster wavelenged either the lelay period a lanthanic	US 1986-831250 CA 1987-529682 JP 1987-34696 US 1986-831250 to an agent contg. a	19870219 19860219 a 1st fluorescent at contg. a rgy emitted, e.g., by E1; E2 then emits do so for a longer cence or of any acce of analyte.

L11 ANSWER 9 OF 10 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER:

1988:91389 HCAPLUS

DOCUMENT NUMBER:

108:91389

TITLE:

Preparation and/or use of 8-hydroxyquinoline

derivatives as enzyme substrates for identification of

microorganisms producing the enzyme

INVENTOR(S):

James, Arthur; Yeoman, Peter

PATENT ASSIGNEE(S):

Cogent Ltd., UK

SOURCE:

Eur. Pat. Appl., 21 pp.

CODEN: EPXXDW

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 238243		19870923	EP 1987-302016	
R: AT, BE, WO 8705632	CH, DE A1	, ES, FR, G 19870924	GR, IT, LI, LU, NL, SE WO 1987-GB165	19870310
W: GB, JP,	US		CD 1086-6032	19860312

PRIORITY APPLN. INFO.:

AB Identification of microorganisms in a sample consists of plating the specimens onto a medium contg. a chromogenic compd. as substrate for the detection of a specific enzyme produced by the microorganism. The enzymic reaction yields a chromogen which is chelated with a metal ion to form a colored ppt. within or around the microorganism. A mixt. of bacteria was multipointed onto agar-peptone medium contg. 8-hydroxyquinoline.beta.-D-galactoside (prepd. from acetobromogalactose and 8-hydroxyquinoline), ferric ammonium citrate, iso-Pr thiogalactoside and read after overnight incubation at 37.degree. Only glucosidase-pos. organisms (e.g. E. coli) produced black colonies. Several hundreds of microorganisms were tested for the enzyme, out of which 80-85% of E. coli in the inoculums were detected but no other enterobacteria gave pos. test results.

L11 ANSWER 10 OF 10 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER:

1987:60780 HCAPLUS

DOCUMENT NUMBER:

106:60780

TITLE:

Rhodium(III) complexes as genotoxic agents:

AUTHOR(S):
CORPORATE SOURCE:
SOURCE:

photochemical effects and their implications LaVelle, James M.; Krause, Ronald A. Sch. Pharm., Univ. Connecticut, Storrs, CT, 06268, USA Mutation Research (1986), 172(3), 211-22 CODEN: MUREAV; ISSN: 0027-5107

DOCUMENT TYPE: Journal

English LANGUAGE: The genetic toxicol. of coordination compds. of transition metals has been of considerable interest since the application of cis-platinum(II) to the therapy of solid tumors. The nature of reactions of such compds. with DNA is still unclear, despite intensive investigation. In this study, several coordination compds. of rhodium(III) were tested for DNA-damaging activity and mutagenicity in bacterial assays in an attempt to understand both the chem. species involved in interactions with DNA and any structural requirements for such interactions. For several complexes, it appears that dissocn. of a ligand from the complex precedes reactions with DNA. This conclusion stems from the finding that photosensitive complexes of rhodium(III) are often many times more toxic to repair-deficient bacterial strains of Escherichia coli K12 when incubated in the light than when incubated in the dark. Similar responses were seen for mutagenicity in Salmonella typhimurium strain TA100. However, reversion of strain TA102 was largely independent of light exposure. Comparisons between mutagenicity and DNA-damaging activity revealed that the 3 activities measured sorted with some independence among the different compds. tested. Thus, the profiles for crosslink formation and (or) generation of oxidative mutagens (mutagenicity in S. typhimurium strain TA102), mutagenicity in TA100 and DNA-damaging activity for the various groups of complexes showed many of the theor. possible combinations of responses in the assays. It is possible, then, that there are different structural requirements for DNA-damaging activity and mutagenicity resp. This may indicate that synthesis of coordination compds. with specific genotoxic properties is possible. Such syntheses may provide complexes for study of DNA-metal interactions and could, later, direct an approach to the design of new antitumor agents.

```
=> d stat que
            197 SEA FILE=REGISTRY ABB=ON (BACTERIOCIN/BI OR BACTERIOCINS/BI)
L1
            114 SEA FILE=REGISTRY ABB=ON NISIN/BI
L2
                                            (LANTIBIOTIC/BI OR LANTIBIOTICS/BI)
             35 SEA FILE=REGISTRY ABB=ON
L3
              1 SEA FILE=REGISTRY ABB=ON COBALT/CN
L4
           4443 SEA FILE=HCAPLUS ABB=ON L1 OR BACTERIOCIN OR ANTIMICROBIAL(W)P
L5
                 EPTIDE? OR BACTERIA? (3W) RIBOSOM? (3W) SYNTH?
           1722 SEA FILE=HCAPLUS ABB=ON L2 OR NISIN 448 SEA FILE=HCAPLUS ABB=ON L3 OR LANTIBIOTIC?
L6
L7
         343310 SEA FILE=HCAPLUS ABB=ON L4 OR COBALT
L8
                                          TRANSITION(W)METAL? OR LANTHANIDE?
         191148 SEA FILE=HCAPLUS ABB=ON
L9
                                           (L8 OR L9)(L)(CHELAT? OR COMPLEX?)
              10 SEA FILE=HCAPLUS ABB=ON
I.11
                 AND (BACTER? OR FUNG?) (W) (DETECT? OR IDEN? OR ASSAY?)
           1790 SEA FILE=HCAPLUS ABB=ON (L5 OR L6 OR L7) AND (?BACTER? OR
L12
                 FUNG?) (L) (DETECT? OR IDEN? OR ASSAY? OR BIND? OR DETERM?)
               1 SEA FILE=HCAPLUS ABB=ON L12 AND (L8 OR L9)
L13
               1 SEA FILE=HCAPLUS ABB=ON L13 NOT L11
L14
```

^{=&}gt; d ibib abs hitrn 114

L14 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER:

1999:622759 HCAPLUS

DOCUMENT NUMBER:

131:334039

TITLE:

Homing in on the role of transition metals in the HNH motif of colicin

endonucleases

AUTHOR(S):

Pommer, Ansgar J.; Kuhlmann, Ulrike C.; Cooper, Alan;

Hemmings, Andrew M.; Moore, Geoffrey R.; James,

Richard; Kleanthous, Colin

CORPORATE SOURCE:

School of Biological Sciences, University of East Anglia, Norwich, NR4 7TJ, UK

SOURCE:

Journal of Biological Chemistry (1999), 274(38),

27153-27160

CODEN: JBCHA3; ISSN: 0021-9258

PUBLISHER:

American Society for Biochemistry and Molecular

Biology

DOCUMENT TYPE:

Journal

English LANGUAGE:

The cytotoxic domain of the bacteriocin colicin E9 (the E9 DNase) is a nonspecific endonuclease that must traverse two membranes to AB reach its cellular target, bacterial DNA. Recent structural studies revealed that the active site of colicin DNases encompasses the HNH motif found in homing endonucleases, and bound within this motif a single transition metal ion (either Zn2+ or Ni2+) the role of which is unknown. In the present work we find that neither Zn2+ nor Ni2+ is required for DNase activity, which instead requires Mg2+ ions, but binding transition metals to the E9 DNase causes subtle changes to both secondary and tertiary structure. Spectroscopic, proteolytic, and calorimetric data show that, accompanying the binding of 1 equiv of Zn2+, Ni2+, or Co2+, the thermodn. stability of the domain increased substantially, and that the equil. dissocn. const. for Zn2+ was less than or equal to nanomolar, while that for Co2+ and Ni2+ was micromolar. Our data demonstrate that the transition metal is not essential for colicin DNase activity but rather serves a structural role. We speculate that the HNH motif has been adapted for use by endonuclease colicins because of its involvement in DNA recognition and because removal of the bound metal ion destabilizes the DNase domain, a likely prerequisite for its translocation

across bacterial membranes. 7440-48-4, Cobalt, biological studies IT

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(binding effect; role of transition metals in the

HNH motif of colicin endonucleases)

REFERENCE COUNT:

THERE ARE 54 CITED REFERENCES AVAILABLE FOR THIS 54 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

```
=> d stat que
                                          (BACTERIOCIN/BI OR BACTERIOCINS/BI)
            197 SEA FILE=REGISTRY ABB=ON
L1
            114 SEA FILE=REGISTRY ABB=ON NISIN/BI
L2
                                           (LANTIBIOTIC/BI OR LANTIBIOTICS/BI)
             35 SEA FILE=REGISTRY ABB=ON
L3
              1 SEA FILE=REGISTRY ABB=ON COBALT/CN
L4
           4443 SEA FILE=HCAPLUS ABB=ON L1 OR BACTERIOCIN OR ANTIMICROBIAL(W)P
L5
                EPTIDE? OR BACTERIA? (3W) RIBOSOM? (3W) SYNTH?
           1722 SEA FILE=HCAPLUS ABB=ON L2 OR NISIN
L6
            448 SEA FILE=HCAPLUS ABB=ON
                                          L3 OR LANTIBIOTIC?
L7
                                          L4 OR COBALT
         343310 SEA FILE=HCAPLUS ABB=ON
\Gamma8
                                          TRANSITION(W) METAL? OR LANTHANIDE?
         191148 SEA FILE=HCAPLUS ABB=ON
L9
                                          (L8 OR L9) (L) (CHELAT? OR COMPLEX?)
             10 SEA FILE=HCAPLUS ABB=ON
L11
                AND (BACTER? OR FUNG?) (W) (DETECT? OR IDEN? OR ASSAY?)
           1790 SEA FILE=HCAPLUS ABB=ON (L5 OR L6 OR L7) AND (?BACTER? OR
I.12
                FUNG?) (L) (DETECT? OR IDEN? OR ASSAY? OR BIND? OR DETERM?)
              1 SEA FILE=HCAPLUS ABB=ON L12 AND (L8 OR L9)
L13
              1 SEA FILE=HCAPLUS ABB=ON L13 NOT L11
L14
              8 SEA L11 OR L14
L15
```

=> d abs bib 115 1-8

L15 ANSWER 1 OF 8 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

The cytotoxic domain of the bacteriocin colicin E9 (the E9 DNase) is a nonspecific endonuclease that must traverse two membranes to reach its cellular target, bacterial DNA. Recent structural studies revealed that the active site of colicin DNases encompasses the HNH motif found in homing endonucleases, and bound within this motif a single transition metal ion (either Zn2+ or Ni2+) the role of which is unknown. In the present work we find that neither Zn2+ nor Ni2+ is required for DNase activity, which instead requires Mg2+ ions, but binding transition metals to the E9 DNase causes subtle changes to both secondary and tertiary structure. Spectroscopic, proteolytic, and calorimetric data show that, accompanying the binding of 1 eq of Zn2+, Ni2+, or Co2+, the thermodynamic stability of the domain increased substantially, and that the equilibrium dissociation constant for Zn2+ was less than or equal to nanomolar, while that for Co2+ and Ni2+ was micromolar. Our datademonstrate that the transition metal is not essential for colicin DNase activity but rather serves a structural role. We speculate that the HNH motif has been adapted for use by endonuclease colicins because of its involvement in DNA recognition and because removal of the bound metal ion destabilizes the DNase domain, a likely prerequisite for its translocation across bacterial membranes.

- AN 1999:482899 BIOSIS
- DN PREV199900482899
- TI Homing in on the role of transition metals in the HNH motif of colicin endonucleases.
- AU Pommer, Ansgar J.; Kuhlmann, Ulrike C.; Cooper, Alan; Hemmings, Andrew M.; Moore, Geoffrey R.; James, Richard; Kleanthous, Colin (1)
- CS (1) School of Biological Sciences, University of East Anglia, Norwich, NR4
- SO Journal of Biological Chemistry, (Sept. 17, 1999) Vol. 274, No. 38, pp. 27153-27160.
 ISSN: 0021-9258.
- DT Article
- LA English
- SL English

- ANSWER 2 OF 8 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC. L15The genetic toxicology of coordination compounds of transition metals has been of considerable interest since the application of cis-platinum(II) to the therapy of solid tumors. The nature of reactions of such compounds with DNA is still unclear, despite intensive investigation. In this study, several coordination compounds of rhodium(III) were tested for DNA-damaging activity and mutagenicity in bacterial assays in an attempt to understand both the chemical species involved in interactions with DNA and any structural requirements for such interactions. For several complexes it appears that dissociation of a ligand from the complex precedes reactions with DNA. This conclusion stems from the finding that photosensitive complexes of rhodium(III) are often many times more toxic to repair-deficient bacterial strains of E. coli K12 when incubated in the light than when incubated in the dark. Similar responses were seen for mutagenicity in S. typhimurium strain TA100. However, reversion of strain TA102 was largely independent of light exposure. Comparisons between mutagenicity and DNA-damaging activity revealed that the 3 activities measured sorted with some independence among the different compounds tests. Thus, the profiles for crosslink formation and/or generation of oxidative mutagens (mutagenicity in S. typhimurium strain TA102), mutagenicity in TA100 and DNA-damaging activity for the various groups of complexes showed many of the theoretically possible combinations of responses in the assays. It is possible, then, that there are different structural requirements for DNA-damaging activity and mutagenicity respectively. This may indicate that synthesis of coordination compounds with specific genotoxic properties is possible. Such syntheses may provide complexes for study of DNA-metal interactions and could, later, direct an approach to the design of new antitumor agents.
- 1987:148370 BIOSIS
- DN
- RHODIUM-III COMPLEXES AS GENOTOXIC AGENTS PHOTOCHEMICAL EFFECTS AND THEIR ΤI IMPLICATIONS.
- LAVELLE J M; KRAUSE R A ΑU
- TOXICOL. PROGRAM, SECT. OF PHARMACOL. AND TOXICOL., SCH. OF PHARMACY, CS UNIV. OF CONN., STORRS, CONN. 06268.
- MUTAT RES, (1986) 172 (3), 211-222. SÓ CODEN: MUREAV. ISSN: 0027-5107.
- BA; OLD FS
- English LA
- ANSWER 3 OF 8 MEDLINE L15
- The cytotoxic domain of the bacteriocin colicin E9 (the E9 DNase) is a nonspecific endonuclease that must traverse two membranes to AΒ reach its cellular target, bacterial DNA. Recent structural studies revealed that the active site of colicin DNases encompasses the HNH motif found in homing endonucleases, and bound within this motif a single transition metal ion (either Zn(2+) or Ni(2+)) the role of which is unknown. In the present work we find that neither Zn(2+) nor Ni(2+) is required for DNase activity, which instead requires Mq(2+) ions, but binding transition metals to the E9 DNase causes subtle changes to both secondary and tertiary structure. Spectroscopic, proteolytic, and calorimetric data show that, accompanying the **binding** of 1 eq of Zn(2+), Ni(2+), or Co(2+), the thermodynamic stability of the domain increased substantially, and that the equilibrium dissociation constant for Zn(2+) was less than or equal to nanomolar, while that for Co(2+) and Ni (2+) was micromolar. Our

data demonstrate that the **transition** metal is not essential for colicin DNase activity but rather serves a structural role. We speculate that the HNH motif has been adapted for use by endonuclease colicins because of its involvement in DNA recognition and because removal of the bound metal ion destabilizes the DNase domain, a likely prerequisite for its translocation across **bacterial** membranes.

AN 1999410457 MEDLINE

DN 99410457 PubMed ID: 10480931

TI Homing in on the role of transition metals in the HNH motif of colicin endonucleases.

AU Pommer A J; Kuhlmann U C; Cooper A; Hemmings A M; Moore G R; James R; Kleanthous C

CS School of Biological Sciences, University of East Anglia, Norwich NR4 7TJ, United Kingdom.

SO JOURNAL OF BIOLOGICAL CHEMISTRY, (1999 Sep 17) 274 (38) 27153-60. Journal code: 2985121R. ISSN: 0021-9258.

CY United States

DT Journal; Article; (JOURNAL ARTICLE)

LA English

FS Priority Journals

EM 199910

ED Entered STN: 19991026

Last Updated on STN: 19991026 Entered Medline: 19991013

L15 ANSWER 4 OF 8 MEDLINE .

- The genetic toxicology of coordination compounds of transition metals has been of considerable interest since the application of cis-platinum(II) to the therapy of solid tumors. The nature of reactions of such compounds with DNA is still unclear, despite intensive investigation. In this study, several coordination compounds of rhodium(III) were tested for DNA-damaging activity and mutagenicity in bacterial assays in an attempt to understand both the chemical species involved in interactions with DNA and any structural requirements for such interactions. For several complexes it appears that dissociation of a ligand from the complex precedes reactions with DNA. This conclusion stems from the finding that photosensitive complexes of rhodium(III) are often many times more toxic to repair-deficient bacterial stains of E. coli K12 when incubated in the light than when incubated in the dark. Similar responses were seen for mutagenicity in S. typhimurium strain TA100. However, reversion of strain TA102 was largely independent of light exposure. Comparisons between mutagenicity and DNA-damaging activity revealed that the 3 activities measured sorted with some independence among the different compounds tested. Thus, the profiles for crosslink formation and/or generation of oxidative mutagens (mutagenicity in S. typhimurium strain TA102), mutagenicity in TA100 and DNA-damaging activity for the various groups of complexes showed many of the theoretically possible combinations of response in the assays. It is possible, then, that there are different structural requirements for DNA-damaging activity and mutagenicity respectively. This may indicate that synthesis of coordination compounds with specific genotoxic properties is possible. Such syntheses may provide complexes for study of DNA-metal interactions and could, later, direct an approach to the design of new antitumor agents.
- AN 87064816 MEDLINE
- DN 87064816 PubMed ID: 3537777
- TI Rhodium(III) complexes as genotoxic agents: photochemical effects and their implications.

Page 13

10 /082,618 Lucas

LaVelle J M; Krause R A ΑU

MUTATION RESEARCH, (1986 Dec) 172 (3) 211-22. SO Journal code: 0400763. ISSN: 0027-5107.

CY Netherlands

Journal; Article; (JOURNAL ARTICLE) DT

English LA

Priority Journals FS

198612 EM

Entered STN: 19900302 ED

Last Updated on STN: 19970203 Entered Medline: 19861231

L15 ANSWER 5 OF 8 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V.

The cytotoxic domain of the bacteriocin colicin E9 (the E9 AB DNase) is a nonspecific endonuclease that must traverse two membranes to reach its cellular target, bacterial DNA. Recent structural studies revealed that the active site of colicin DNases encompasses the HNH motif found in homing endonucleases, and bound within this motif a single transition metal ion (either Zn2+ or Ni2+) the role of which is unknown. In the present work we find that neither Zn2+ nor Ni2+ is required for DNase activity, which instead requires Mg2+ ions, but binding transition metals to the E9 DNase causes subtle changes to both secondary and tertiary structure. Spectroscopic, proteolytic, and calorimetric data show that, accompanying the binding of 1 eq of Zn2+, Ni2+, or Co2+, the thermodynamic stability of the domain increased substantially, and that the equilibrium dissociation constant for Zn2+ was less than or equal to nanomolar, while that for Co2+ and Ni2+ was micromolar. Our data demonstrate that the transition metal is not essential for colicin DNase activity but rather serves a structural role. We speculate that the HNH motif has been adapted for use by endonuclease colicins because of its involvement in DNA recognition and because removal of the bound metal ion destabilizes the DNase domain, a likely prerequisite for its translocation across bacterial membranes.

1999326741 EMBASE AN

Homing in on the role of transition metals in the HNH ΤI motif of colicin endonucleases.

Pommer A.J.; Kuhlmann U.C.; Cooper A.; Hemmings A.M.; Moore G.R.; James ΑU R.; Kleanthous C.

C. Kleanthous, School of Biological Sciences, University of East Anglia, CS Norwich NR4 7TJ, United Kingdom. c.kleanthous@uea.ac.uk

Journal of Biological Chemistry, (17 Sep 1999) 274/38 (27153-27160). SO Refs: 54 ISSN: 0021-9258 CODEN: JBCHA3

United States CY

Journal; Article DT

Microbiology FS 004

English LA

English SL

ANSWER 6 OF 8 JICST-EPlus COPYRIGHT 2003 JST

DV-7572 injection was examined for mutagenic activity in the reversion AΒ test with bacterial assays using Salmonella typhimurium strains TA1535, TA1537, TA98 and TA100 and Escherichia coli strain, WP2uvrA. The doses for the assays were selected from a dose rangefinding study conducted on the test material at dose levels of 1.23, 2.45, 4.90, 9.81, 19.6, 39.2, 78.5, 157, 314, 628, 1256, 2511, 5022 and 10045 .MU.g per plate using the S.typhimurium strain, TA100 and the E.coli strain WP2uvrA. The test material was nontoxic to both strains in this

dose rangefinding study. The dose selected for the mutation assays were 1,10,100,500,1000,2499,4998 and9997 .MU.g per plate for the independent repeat assays. The test material, DV-7572 injection did not exhibit genetic activity in these assays and was not mutagenic under these test conditions according to our assay criteria. (author abst.)

930550957 JICST-EPlus AN

- Mutagenicity Studies of DV-7572 Injection (1): Reversion Test with ΤI Bacteria.
- JAGANNATH D R SHIMADA HIROYASU
- Hazleton Washington Inc. CS Daiichi Seiyaku Co., Ltd.
- Yakuri to Chiryo (Japanese Pharmacology & Therapeutics), (1993) vol. 21, SO no. Suppl 3, pp. S.875-S.879. Journal Code: Z0947A (Tbl. 4, Ref. 4) ISSN: 0386-3603
- Japan CY
- Journal; Article DT
- Japanese LA
- New STA
- L15 ANSWER 7 OF 8 JICST-EPlus COPYRIGHT 2003 JST
- To determine the value of repeated intraperitoneal cisplatin(CDDP) AB administration in treating malignant ovarian tumor(ip treatment), we studied the intracorporeal dynamics and tissue concentrations of CDDP as well as its safety and clinical effects in 29 cases. The blood concentration of total-Pt showed higher preadministration levels with the increase in the number of administrations, The difference continued to increase at the same rate for 24 to 72 hours after administration. On the other hand, free-Pt was not detected in the blood before each administration even after repeated administrations, but showed relatively high levels immediately after the ip treatment up to the 2nd hour. Also the changes in free-Pt concentration in the ascites showed a pattern similar to that in the blood. On initial administration of CDDP, the total-Pt concentration in the overian tumor tissue was 0.19-0.76.MU.g/g in iv. 1.26-1.84 .MU.g/g in ip, and 0.89-4.85.MU.g/g in ia. The total-Pt concentration in the ovarian tumor tissue following repeated ip treatments tended to be higher than that after the first ip treatment. After 10 or more repeated ip tretment, no tendency of aggravation in renal function was observed. The bacterial detection rate by an indweling intraperioneal tube was 40.5 %(15/47), and almost all isolated bacteria were CNS. A slight increase in the WBC was recognized in 2 cases (abridged author abst.)

930036956 JICST-EPlus ΑN

- Intracorporeal dynamics, safety and clinical results of repeated TΙ intraperitoneal cisplatin (CDDP) administration in treating malignant ovarian tumors.
- IWASA TAKESHI; USUI NAOYUKI; SUZUKI MASAAKI; TAKADA MICHIO ΑU
- Juntendo Univ., School of Medicine CS
- Juntendo Igaku (Juntendo Medical Journal), (1992) vol. 38, no. 3, pp. SO 418-427. Journal Code: G0715A (Fig. 9, Tbl. 3, Ref. 19) CODEN: JUIZAG; ISSN: 0022-6769
- CYJapan
- Journal; Article DT
- Japanese LA
- New STA
- ANSWER 8 OF 8 WPIDS (C) 2003 THOMSON DERWENT L15
- 2001-281830 [29] WPIDS AN
- 2001-282097 [29] CR

WO 200126673 A UPAB: 20010822 AΒ NOVELTY - A complex (I) comprising a cyclic antibiotic and at least one of a lanthanide or a transition metal is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for:

- (1) (I) comprising polymyxin (especially polymyxin B or colistin) and a metal;
- (2) detecting gram negative bacteria in a sample suspected of containing gram negative bacteria, comprises contacting the sample with (I) such that the complex binds to the gram negative bacteria to yield a bound complex, separating the bound complex from any nonbound complex, where the presence of a bound complex is indicative of the presence of gram negative bacteria;
- (3) detecting disease in a patient suspected of having the disease, comprising introducing a detectable complex comprising a cyclic antibiotic, a metal and a delivery molecule into the patient, where the delivery molecule targets the complex to a disease cell, if present, and detecting the presence or absence of the complex at a site within the patient, where the presence of the complex at the site is indicative of the presence of a disease in the patient site;
- (4) detecting the presence of gram negative bacteria in a patient suspected of comprising gram negative bacteria, comprising introducing a detectable complex containing a cyclic antibiotic and a metal into the patient, and detecting the presence of the complex at the site is indicative of the presence of gram negative bacteria in the patient;
- (5) introducing a detectable complex into a patient, comprising a cyclic antibiotic, a metal and a delivery molecule targeting the complex to a disease cell, to detect disease by detecting the complex at a site, indicative of a disease cell, or treat infection, disease or autoimmune dysfunction; and
- (6) detecting gram negative bacteria in a food sample, comprising incubating the sample with immunomagnetic beads coated with antibody to the gram negative bacterium such that gram negative bacteria bind to the immunomagnetic beads, magnetically removing the immunomagnetic beads from the sample and contacting the immunomagnetic beads with the detectable complex to yield a detectable bound complex, and assaying the immunomagnetic beads for the presence or absence of detectable bound complex, where the presence of a detectable bound complex is indicative of the presence of gram negative bacteria in the food sample.

ACTIVITY - antibacterial; antiautoimmune; cytostatic.

MECHANISM OF ACTION - No details provided.

USE - The complex is useful for detecting gram negative bacteria in samples, especially in food samples, medical samples (e.g. medical fluid) or biological samples (e.g. body tissue), e.g. in food processing or medical sterilization. It is useful to detect gram negative bacteria in patients, by introducing a detectable complex (especially comprising polymyxin B) and detecting the complex at a site within the patient; the complex may also be used therapeutically to kill or disable the gram negative bacteria detected at the site. It may be combined with a delivery molecule e.g. a monoclonal antibody to target the complex to a disease cell (e.g. a bacterial cell, cancer cell or cell involved in autoimmune dysfunction) in a patient, useful diagnostically and therapeutically to detect and treat infection, disease or autoimmune dysfunction (all claimed). Polymyxin B pentasulfate (80 mg, 0.05 mmol) was dissolved in 5 ml 0.05 M acetate buffer, pH 5.5, incubated at room temperature with cobalt chloride (12 mg, 0.055 mmol) and purified by column

Lucas

chromatography by known methods. UV-absorbing fractions (polymyxin B-Cobalt (II) complex) were collected and freeze dried. A titration curve for E. coli O157:H7 was then produced. Bacteria were diluted in sterile saline to 10 CFU (colony forming unit)/ml, incubated (20 minutes room temperature) with 20 micro g/ml polymyxin B-Cobalt (II) complex, centrifuged and resuspended in 0.1 ml saline. Chemiluminescence was measured using 0.2 ml proprietary reagent in a luminometer. A ground beef sample was then tested for E. coli 0157:H7 using a known immunomagnetic capture technique for separation of bacteria from ground beef samples (Pyle et al., Appl. Environ. Microbiol., 65:1966-1972 (1999)), and treatment of collected beads bearing E. coli O157:H7 cells (resuspended in 1.0 ml saline) with 20 micro g/ml polymyxin B-Cobalt (II) complex. Cells were collected in a particle concentrator, re-suspended in 0.1 ml saline and assayed for chemiluminescence, no results are included.

Dwg.0/8

WPIDS AN 2001-281830 [29]

CR 2001-282097 [29]

DNN N2001-200922 DNC C2001-085763

New complex comprising a cyclic antibiotic and a lanthanide or transition metal, useful e.g. for detecting gram negative bacteria in food, medical or biological samples or in diagnosis and treatment of diseases e.g. cancer in patients.

B04 C06 D13 D16 K08 P31 S03 DC

IN FEIRTAG, J M; OLSTEIN, A D

PΑ (KALL-N) KALLESTAD LAB INC

CYC

WO 2001026673 A1 20010419 (200129)* EN 35p PI

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG UZ VN YU ZA ZW

AU 2001010835 A 20010423 (200147)

WO 2001026673 A1 WO 2000-US28358 20001013; AU 2001010835 A AU 2001-10835 20001013

AU 2001010835 A Based on WO 200126673

PRAI US 1999-159142P 19991013